F. ...

3

Ţ



## WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)			
(51) International Patent Classification 6:			(11) International Publication Number: WO 99/57130
C07H 21/04, C12N 15/00, C12Q 1/02, 1/68		A1	(43) International Publication Date: 11 November 1999 (11.11.99)
			CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,
(30) Priority Data: 60/084,329 5 May 1998 (05.05)	•	US	Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of
(71) Applicant (for all designated States except US): GENE LOGIC,			, amendments.

(72) Inventors; and

MD 21046 (US).

(75) Inventors/Applicants (for US only): PRASHAR, Yatindra [IN/US]; 9749 Clocktower Lane #304, Columbia, MD 21046 (US). WEISSMAN, Sherman [US/US]; 459 St. Ronan Street, New Haven, CT 06511 (US).

INC. [US/US]; 708 Quince Orchard Road, Gaithersburg,

- (74) Agent: ADLER, Reid, G.; Morgan, Lewis & Bockius LLP, 1800 M Street, N.W., Washington, DC 20036 (US).
- (54) Title: A PROCESS TO STUDY CHANGES IN GENE EXPRESSION IN T LYMPHOCYTES

## (57) Abstract

Methods are disclosed to identify T lymphocyte genes that are differentially expressed upon exposure to a pathogen (viral or bacterial), immunogen, antigen, or in a sterile inflammatory disease, autoimmune disease, immunodeficiency disease, lymphocytic cancers, or graft versus host rejection. The method involves the preparation of a gene expression profile of a T lymphocyte population exposed to a pathogen or isolated from a subject having one of the aforementioned pathologies and comparing that profile to a profile prepared from quiescent T lymphocytes. The present invention is particularly useful for identifying cytokine genes, genes encoding cell surface receptors and genes encoding intermediary signalling molecules. Related methods for identifying therapeutic or prophylactic immunomodulatory agents are presented. Articles of manufacture are disclosed that comprise selected grouping of nucleic acids, affixed to a solid support, that correspond to genes that are differentially expressed in various populations or subpopulations of T lymphocytes at variations stages of T cell differentiation, in quiescent versus activated T lymphocytes or normal versus diseased T lymphocytes.